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- (9) Cross-linked polymers for contact lenses.
- (57) Contact lenses are disclosed which are formed from hydrophilic polymers of at least one hydrophilic monomer and minor proportions of at least one monomer containing cationic groups and at least one monomer containing anionic groups so that ionic cross-linkages can form between polymer molecules. Typical monomers containing cationic groups are alkyl amino acrylates and methacrylates and typical monomers containing anionic groups are methacrylic and acrylic acids and such monomers may be employed in proportions by weight of from about 5% to about 20%. Suitable hydrophilic monomers include hydroxy alkyl acrylates and methacrylates.

CROSS-LIMKED POLYMERS FOR CONTACT LEMSES

The present invention relates to hydrophilis polymers having ionis cross-linkages, contact lenses formed therefrom and a process for their preparation.

5 BACKGROUND OF THE INVENTION

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Cross-linking of polymers is well-known in the art for providing polymers with desirable properties, in particular desirable mechanical properties such as high tensile strength. Cross-linking is usually effected by 10 use of a covalent cross-linking agent which bridges two or more polymer chains linking them together by covalent bonding. Cross-linking can also be effected by means of electrovalent interactions involving ionized sites in neighbouring polymer chains; cross-links induced in this way are known as ionic cross-links. This form of cross-15 linking is illustrated by ionomer resins which are ethylene copolymers containing pendant carboxylic acid groups and an associated bivalent metal cation and also by polyelectrolyte complexes which consist of an essentially neutralized 20 mixture of polycations (e.g. a polyquaternary ammonium compound and a polyanion such as poly (vinyl sulfonia acid).

Cross-linking of hydrophilic polymers for use as contact lens materials is often a practical necessity in order that the material has adequate mechanical properties in the hydrated state. Covalent cross-linkage has been widely used for this purpose but ionic cross-linking has not hitherto been used. Ionic cross-linking requiring the

presence of a bivalent metal cation is not acceptable for contact lens material since in use the bivalent cations are largely leacned from the polymer by tear fluid and replaced by sodium and potassium ions contained in the tear fluid; polyelectrolyte complexes are unsuitable since they cannot conveniently be prepared in a form suitable for machining on a lathe.

BRIEF DESCRIPTION OF THE INVENTION

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According to one aspect of the invention, there

is provided a novel polymer for use as a contact lens
material and comprised of chains in each of which there
are both cationic and anionic sites participating in
electrovalent cross-linkage of the chains one to another,
the aforesaid sites being provided by moieties of the polymer

chains which are resistant to loss of polarity by contact
with tear fluids.

In a more specific embodiment the invention provides a contact lens which comprises a hydrophilic polymer of at least one hydrophilic vinyl or vinylidene monomer, a minor proportion of a cationic group + containing monomer and a minor proportion of an anionic group - containing monomer and said ionic group - containing monomers providing cationic and anionic sites for ionic cross-linkages between polymer chains.

The moieties providing the ionic sites will usually be organic moieties polymerized into the polymer chains and in general such organic moieties will be units

of monomer material copolymerized into a backbone polymer. The monomer material may, for example, be an alkyl amino acrylate or methacrylate, such as dimethylaminoethylmethacrylate, t-butylaminoethylmethacrylate or di-n-butylaminoethylmethacrylate, in the case of monomer material 5 providing cationic sites. Conveniently methacrylic or acrylic acid may be employed to provide anionic sites although other polymerizable carboxylic acids may be used. Alternatively, however, such other cationic and anionic moieties as quaternary ammonium compounds 10 and unsaturated sulfonic acids may be employed to provide the cationic and amionic sites although, it should be noted, particularly in the case of relatively strongly ionizing groups, that there may be difficulties as regards availability of materials. Vinyl sulfonic acid, styrene 15 sulfonic acid and unsaturated phosphoric acid monomers are however, obtainable.

The polymers of the invention will normally contain the ionic moieties in a comparatively minor proportion together with a balance of polymerized essentially non-ionic monomer material. The balance may, for example, be made up of polymerized hydrophilic monomer such as an hydroxy alkyl acrylate or methacrylate, e.g. 2-hydroxy-ethylmethacrylate or N-vinylpyrrolidone and/or a polymerized hydrophobic monomer, such as an alkyl acrylate or methacrylate or styrene or a substituted styrene. Typical hydrophobic monomers are methyl methacrylate and n-butyl methacrylate.

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Copplymens of O-nyaroxyethylmethacrylate with nimethylaminoethylmethacrylate, methacrylic acid and optionally styrene have seen found to be especially suitable for use as soft contact lens material.

The anionic moreties may be present in substantially stoichiometrically equivalent amounts, but, if desired, one ionic moiety may be in stoichiometric excess. For example, a stoichiometric excess of the anionic moiety provides a powerful hydration mechanism in the alkaline environment of the eye so that in most instances such an excess will be considered desirable. In such cases, where the ionic groups are in excess they are preferably neutralised by treatment of the final polymer with a mildly alkaline solution as described in British Patent No. 1,548,158

(the disclosure of which is specifically incorporated herein by reference).

As indicated earlier, the ionic moieties will in general be present in relatively minor proportions. Generally the proportion of ionic moieties available for participation in electrovalent linkage will be in the range from about 5% to about 20% by weight, preferably 7% to 20% (e.g. from about 7% to 17% by weight). In all cases, the percentages stated are by weight of the whole unhydrated polymer.

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Polymers according to the invention can conveniently be made by polymerizing a monomer mixture comprising components of identity and content appropriate to the desired polymer, the polymerization being effected using, for example, azobisisobutyronitrile as initiator. Other initiators may alternatively be employed although peroxile initiators are not at present recommended since they have been found to produce dark poloured polymers.

The polymers of the invention have been found.

5 as will be appreciated from the Examples which follow, to exhibit good mechanical properties. This is thought to be a result of the labile character of the electrovalent cross-linkage forces which results in the polymeric material being able to yield locally to applied stress. The polymers of the invention may, in addition to cross-linkages as a result of electrovalency, contain covalent cross-linkages as a result of use, if desired, of a covalent cross-linking agent, and results in polymers having even better mechanical properties. Covalent cross-linkages may be incorporated in the polymers by including a di- or poly functional cross-linking agent, such as glycol dimethacrylates or acrylates, allyl methacrylate or acrylate or triallyl cyanurate.

The polymer of the invention can readily be cast to form a lens or lens blank. Lens blanks can be macrined

- 20 in known manner to form contact lenses of desired configuration.

 Lenses made in this way from hydrophilic monomer-derived polymers can be hydrated to equilibrium after machining using saline solution as described, for example, in British Patent No. 1,548,158.
- The invention includes within its scope lenses, blanks and machined lenses as just described and in particular includes such lenses and lens blanks made of polymer

comprising the following polymerized monomer units in the following proportions by weight:-

2-hydroxyethylmethacrylate: 24 parts (2-HEMA)

dimethylaminoethylmethacrylate: from 1 to 2.25 parts (DMAEMA)

methacrylic acid: from 1 to 3 parts (MA)

styrene: .

up to 2.5 parts

The following specific Examples are intended to illustrate the invention.

Examples 1 to 5

5

Polymers were prepared according to the formulations 15 given in Table 1. 2-hydroxyethylmethacrylate (2-HEMA) with a very low (0.05 wt%) ethylene glycol dimethacrylate (EGDM) content was used and hence the polymer was expected to contain a very low concentration of derived covalent crosslinks. The monomer mixtures were polymerized by thoroughly 20 mixing together purified 2-HEMA, redistilled MA, DMAEMA and (where stated styrene) in the proportions stated in Table 1 using the stated quantity of azodiisobutyronitrile (AZDN) as free radical generator. Polymerization was carried out in polythene tubes at 65°C for 16 hours, followed by curing 25 for 3 hours at 100°C. In the case of comparison Examples 1 and 3, the DMAEMA was omitted, as indicated.

The rods obtained were cut into lens blanks, machined to form lenses, extracted and hydrated to equilibrium in a saline solution buffered at pH7.4. The

pronounce encourage is arrived and the recalls are provided in Tubile 2. It can be seen that polymers of augustications surranged as judged by the curbt test, can be propared even in the apparent assence of covalent prosecutions.

This observation and the fact that the water content of the gels containing DMAEMA are somewhat lower than those without, in spite of the hydrophilic nature of DMAEMA and twice the addition of methacrylic acid, provides convincing evidence that hydrogels which contain minor proportions of cationic and anionic sites contain physically—effective cross-links attributable to the simultaneous presence of both types of ionic site.

TABLE 1.

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20

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| | 1 | 2 | 3 | 4 | 5 |
|-----------------------|-------|-------|-------|-------|-------------|
| 2-HIMA (purified) (g) | 24.30 | 24.00 | 24.00 | 24.00 | 24.00 |
| DMAEMA (g) | 0.00 | 1.23 | 0.00 | 2.00 | i : 2.00 |
| Methacrylic Acid (g) | 0.68 | 1.20* | 1.77 | 2.09* | 2.35 |
| Styrene (g) | 0.00 | 0.00 | 2.05 | 2.05 | 2.35 |
| AZUN (g) | 0.02 | 0.02 | 0.03 | 0.03 | ! : 0.03 |

* stoichiemetric excess with respect to the amount of dimethylaminoethylmethacrylate (DMADMA)

TABLE 2

| | Example No. | Extractable Concent (% w/w) | Mater Content (3 W/W) | S#G11 3) | Burst Prosoure Dens Thiokness |
|----|-------------------|-----------------------------|-----------------------------|-------------|--|
| 5 | 1 (comparison) | 5.3 | 64 | ÷ 5 | 3 psi, 0.21 5, |
| | 2 | 7.9 | 59 | 35 | 16 psi/3.83mm |
| 10 | 3 (comparison) | 5.5 | 71 | 55 | 3 psi/0.10mm |
| | 4 | 11.5 | 55 | 32 | 19 psi/0.20 m. |
| | 5 | 8.8 | 64 | 42 | 15 psi/0.llmm |

<u>CLAIMS</u>

- A polymeric contact lens formed by addition polymerization of at least one hydrophilis, ethylenicallyunsaturated monomer, an ethylenically-unsaturated monomer containing a cationic group and an ethylenically-
- 5 unsaturated monomer containing an anionic group, said cationic and anionic groups forming ionic cross-linkages whereby the lens exhibits improved burst strength as compared with a similar lens containing no ionic crosslinkages.
- 10 2. A contact lens according to claim 1 wherein the monomer containing a cationic group is a vinyl or vinylidene monomer having an amino group or quaternary ammonium group.
- 3. A contact lens according to claim 2 wherein the 15 vinyl or vinylidene monomer is an alkyl amino acrylate or
 - methacrylate.
- A contact lens according to any one of the preceding claims wherein the monomer containing an anionic group is a vinyl or vinylidene monomer having a 20 carboxylic or sulfonic acid group.
 - A contact lens which comprises a polymer of a hydrophilic vinyl or vinylidene monomer containing a single center of unsaturation, a cationic vinyl or vinylidene monomer having alkyl amino or quaternary

ammonium dationic groups and an anionic vinyl or vinylidene monomer naving carboxylic or sulfonic acid anionic groups, said dationic and anionic groups constituting pendant or terminal groups in the polymer which interact to form ionic cross-linkages.

- 5 8. A contact lens according to claim 5 wherein the anionic monomer is present in the polymer in stoichiometric excess with respect to the cationic monomer.
 - 7. A contact lens according to claim 5 or claim 8 wherein the cationic and anionic monomers are present
- 10 in the polymer in a total amount of from about 5 to about 20% by weight.
 - 8. A contact lens according to any one of claims
 5 to 7 wherein the polymer also contains covalent crosslinkages derived from cross-linking monomers containing
- 15 at least two centers of unsaturation.
 - 9. A contact lens according to any one of claims
 5 to 8 wherein the polymer is an addition polymer of at
 least one hydrophilic monomer and at least one hydrophopic
 monomer.
- 20 10. A contact lens which comprises a polymeric hydrogel which is the hydrated polymerized product of a monomer mixture comprising a minor proportion by weight of vinyl or vinylidene monomers containing cationic and anionic groups and a major proportion of polymerizable
- monomer having a single center of unsaturation, said monomer mixture including sufficient hydrophilic monomer to render the resultant polymer hydrophilic and the

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- (34) Cross-linked polymers for contact lenses.
- © Contact lenses are disclosed which are formed from hydrophilic polymers of at least one hydrophilic monomer and minor proportions of at least one monomer containing cationic groups and at least one monomer containing anionic groups so that ionic cross-linkages can form between polymer molecules. Typical monomers containing cationic groups are alkyl amino acrylates and methacrylates and typical monomers containing anionic groups are methacrylic and acrylic acids and such monomers may be employed in propartions by weight of from about 5% to about 20%. Suitable hydrophilic monomers include hydroxy alkyl acrylates and methacrylates.

monoment containing dations and anionic groups being effective in comming conto pross-linkages in the resultant polymen.

- addition polymer of at least one hydrophilis vinyl or vinylidene monomer. A minor proportion by weight of a copolymerizable monomer containing anionic groups and a minor proportion by weight of a copolymerizable monomer containing cationic groups, the anionic and cationic groups forming ionic cross-linkages.
 - A method of preparing a hydrophilic copolymer containing ionic cross-linkages which comprises subjecting to polymerization in the presence of a free-radical initiator a monomer mixture comprising at least one
- hydrophilic vinyl or vinylidene monomer containing a single center of unsaturation, a vinyl or vinylidene monomer containing a cationic group and a vinyl or vinylidene monomer containing anionic groups.

Application number

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| | | | | CATEGORY OF CITED DOCUMENTS X: particularly relevant A: technological background O: non-written disclosure P: intermediate document T: theory or principle underlying |
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